```
SVELD, Jeroen
        AT, Sebastianus Antonius Johannes
      DANKERT, Jacob
      KUIJPERS, Alma Johanna
      ENGBERS, Gerardus Henricus Maria
                                                            RECEIVED
      FEIJEN, Jan
<120> Isolated and Recombinant Antimicrobial Peptides
                                                         TECH CENTER 1600/2900
      Thrombocidin-1 (TC-1) and Thrombocidin-2 (TC-2)
      or Variants Thereof
<130> 702 000648
<140> 09/509,391
<141> 2000-07-07
<150> EP 97202934.2
<151> 1997-09-25
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<150> EP 98201411.0
                                                     APR 1 7 2002
<151> 1998-05-01
                                                 TECH CENTER 1600/2900
<160> 18
<170> MS Word 97 SR-2
<210> 1
<211> 85
<212> PRT
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<223> CTAP-III (connective tissue activating peptide)
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Glu Leu Arg Cys Met Cys Ile Lys Thr Thr Ser Gly Ile His Pro Lys
Asn Ile Gln Ser Leu Glu Val Ile Gly Lys Gly Thr His Cys Asn Gln
Val Glu Val Ile Ala Thr Leu Lys Asp Gly Arg Lys Ile Cys Leu Asp
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<223> N-terminus of TC-1a and TC-1b thrombocidins

<400> 2

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Ala Glu Leu Arg

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<223> TC-1* thrombocidin variant
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Ala Glu Leu Arg Cys Met Cys Ile Lys Thr Thr Ser Gly Ile His Pro
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Lys Asn Ile Gln Ser Leu Glu Val Ile Gly Lys Gly Thr His Cys Asn
                                 25
Gln Val Glu Val Ile Ala Thr Leu Lys Asp Gly Arg Lys Ile Cys Leu
                             40
Asp Pro Asp Ala Pro Arg Ile Lys Lys Ile Val Gln Lys Lys Leu Ala
Gly Asp Glu Ser
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<223> N-terminus of TC-1d thrombocidin
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Tyr Ala Glu Leu Arg
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Ala Gly Asp Glu Ser
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<211> 83
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<223> TC-2 thrombocidin
<400> 6
Asn Leu Ala Lys Gly Lys Glu Glu Ser Leu Asp Ser Asp Leu Tyr Ala
                                     10
Glu Leu Arg Cys Met Cys Ile Lys Thr Thr Ser Gly Ile His Pro Lys
Asn Ile Gln Ser Leu Glu Val Ile Gly Lys Gly Thr His Cys Asn Gln
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Val Glu Val Ile Ala Thr Leu Lys Asp Gly Arg Lys Ile Cys Leu Asp

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50
Pro Asp Ala Pro Arg Ile Lys Lys Ile Val Gln Lys Lys Leu Ala Gly
                                           75
Asp Glu Ser
<210> 7
<211> 34
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<223> Forward primer
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<221> misc_feature
<222> (5)...(10)
<223> BamHI restriction site
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tataggatcc atgagcctca gacttgatac cacc
                                                                        34
<210> 8
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<222> (11) ... (13)
<223> Stop sequence
<400> 8
tataggatcc tcaatcagca gattcatcac ctgccaat
                                                                        38
<210> 9
<211> 33
<212> DNA
<213> Homo sapiens
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<223> Forward primer for CTAP-III and TC-2
<220>
<221> misc_feature
<222> (7)...(12)
<223> NdeI restriction site
<400> 9
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gtgtaacata tgaacttggc gaaaggcaaa gag
<210> 10
<211> 33
<212> DNA
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<213> Homo sapiens
<220>
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<210> 11
<211> 36
<212> DNA
<213> Homo sapiens
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<223> Forward primer for TC-1
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<221> misc_feature
<222> (7)...(12)
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                                                                       36
<210> 12
<211> 68
<212> PRT
<213> Homo sapiens
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<223> TC-1 thrombocidin
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Leu Arg Cys Met Cys Ile Lys Thr Thr Ser Gly Ile His Pro Lys Asn
                                                          15
Ile Gln Ser Leu Glu Val Ile Gly Lys Gly Thr His Cys Asn Gln Val
             20
                                 25
Glu Val Ile Ala Thr Leu Lys Asp Gly Arg Lys Ile Cys Leu Asp Pro
Asp Ala Pro Arg Ile Lys Lys Ile Val Gln Lys Lys Leu Ala Gly Asp
    50
Glu Ser Ala Asp
65
<210> 13
<211> 70
<212> PRT
<213> Homo sapiens
<223> NAP-2 (neutrophil activating peptide)
<400> 13
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Lys Asn Ile Gln Ser Leu Glu Val Ile Gly Lys Gly Thr His Cys Asn
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20
Gln Val Glu Val Ile Ala Thr Leu Lys Asp Gly Arg Lys Ile Cys Leu
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Asp Pro Asp Ala Pro Arg Ile Lys Lys Ile Val Gln Lys Lys Leu Ala
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Gly Asp Glu Ser Ala Asp
<210> 14
<211> 69
<212> PRT
<213> Escherichia coli
<223> rMTC-1* (TC-1* carrying an additional N-terminal methionine)
Met Ala Glu Leu Arg Cys Met Cys Ile Lys Thr Thr Ser Gly Ile His
Pro Lys Asn Ile Gln Ser Leu Glu Val Ile Gly Lys Gly Thr His Cys
                                  25
Asn Gln Val Glu Val Ile Ala Thr Leu Lys Asp Gly Arg Lys Ile Cys
                             40
Leu Asp Pro Asp Ala Pro Arg Ile Lys Lys Ile Val Gln Lys Lys Leu
                         55
Ala Gly Asp Glu Ser
65
<210> 15
<211> 84
<212> PRT
<213> Escherichia coli
<220>
<223> rMTC-2 (TC-2 carrying an additional N-terminal methionine)
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Ala Glu Leu Arg Cys Met Cys Ile Lys Thr Thr Ser Gly Ile His Pro
             20
                                 25
Lys Asn Ile Gln Ser Leu Glu Val Ile Gly Lys Gly Thr His Cys Asn
                             40
Gln Val Glu Val Ile Ala Thr Leu Lys Asp Gly Arg Lys Ile Cys Leu
    50
                         55
Asp Pro Asp Ala Pro Arg Ile Lys Lys Ile Val Gln Lys Lys Leu Ala
                     70
Gly Asp Glu Ser
<210> 16
<211> 91
<212> PRT
<213> Escherichia coli
<223> rYTC-1 (TC-1 with an N-terminal His-tag, plus a tyrosine residue)
<220>
<221> SITE
<222> (1)...(21)
<223> Antimicrobial activity enhancing sequence (Histag)
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<400> 16 Met Gly His His His His His His His His His Ser Ser Gly His 10 Ile Glu Gly Arg His Met Tyr Leu Arg Cys Met Cys Ile Lys Thr Thr 25 Ser Gly Ile His Pro Lys Asn Ile Gln Ser Leu Glu Val Ile Gly Lys 40 45 Gly Thr His Cys Asn Gln Val Glu Val Ile Ala Thr Leu Lys Asp Gly 55 60 Arg Lys Ile Cys Leu Asp Pro Asp Ala Pro Arg Ile Lys Lys Ile Val 70 Gln Lys Lys Leu Ala Gly Asp Glu Ser Ala Asp <210> 17 <211> 93 <212> PRT <213> Escherichia coli <223> rYNAP (NAP with an N-terminal His-tag, plus a tyrosine residue) <220> <221> SITE <222> (1)...(21) <223> Antimicrobial activity enhancing sequence (Histag) Met Gly His His His His His His His His Ser Ser Gly His 10 Ile Glu Gly Arg His Met Tyr Ala Glu Leu Arg Cys Met Cys Ile Lys Thr Thr Ser Gly Ile His Pro Lys Asn Ile Gln Ser Leu Glu Val Ile Gly Lys Gly Thr His Cys Asn Gln Val Glu Val Ile Ala Thr Leu Lys Asp Gly Arg Lys Ile Cys Leu Asp Pro Asp Ala Pro Arg Ile Lys Lys Ile Val Gln Lys Lys Leu Ala Gly Asp Glu Ser Ala Ile <210> 18 <211> 86 <212> PRT <213> Escherichia coli <220> <223> rMCTAP (CTAP with an additional N-terminal methionine) <400> 18 Met Asn Leu Ala Lys Gly Lys Glu Glu Ser Leu Asp Ser Asp Leu Tyr 10 Ala Glu Leu Arg Cys Met Cys Ile Lys Thr Thr Ser Gly Ile His Pro 25 Lys Asn Ile Gln Ser Leu Glu Val Ile Gly Lys Gly Thr His Cys Asn Gln Val Glu Val Ile Ala Thr Leu Lys Asp Gly Arg Lys Ile Cys Leu Asp Pro Asp Ala Pro Arg Ile Lys Lys Ile Val Gln Lys Lys Leu Ala

.3

Gly Asp Glu Ser Ala Asp